REPORT DOCUMENTATION PAGE

Form Approved OMB NO. 0704-0188

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4. TITLE A	ND SUBTITLE			5a. C	5a. CONTRACT NUMBER			
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6. AUTHOR	S			5d. P	ROJE	CT NUMBER		
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P.O. Box 12		25500 2211			NUMBER(S)			
Research Ti	riangle Park, NC	27709-2211			62226-LS-DRP.4			
12. DISTRIE	SUTION AVAIL	IBILITY STATI	EMENT					
Approved for	Public Release;	Distribution Un	limited					
	MENTARY NO							
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Report Title

Using Estimations of Entropy to Optimize Complex Human Dynamic Networks under Stress Final Report

ABSTRACT

For all team activities, whether it be sports teams or military units, winning is the ultimate goal. To truly achieve team success, the team should be able to complete a task with the least amount of moves or use the least amount of energy without sacrificing quality.

In this study, we are trying to determine the factors that contribute the most to successful teams under stressful conditions. We hypothesize that by measuring individual biological responses, such as heart rate, breathing rate, posture, temperature, and saliva/blood components, during various physically and mentally stressful exercises, we will be able to determine the factors that drive overall team success and assemble more effective teams using these factors. Sixteen WVU Air Force ROTC participants were selected, divided into four different teams of four individuals, their biological responses were monitored (some in real-time and some prior to and immediately following) in response to stressful teamwork exercises (mock hostage rescue). Individuals were outfitted with EEG, heart rate, breathing rate, estimated core temperature, sound, activity and posture monitors and tasked with finding the "hostage" and moving it to a safe location; to move the hostage required the assembly of a make-shift gurney from items hidden in an urban setting. These exercises were expected to simulate aspects of the types of stresses endured by U.S. Military Special Operations Forces and Hostage Rescue Teams of the Federal Bureau of Investigation.

Enter List of papers submitted or published that acknowledge ARO support from the start of the project to the date of this printing. List the papers, including journal references, in the following categories:

(a) Papers published in peer-reviewed journals (N/A for none)

Received Paper

02/27/2014 3.00 Holly N. Currie, Julie A. Vrana, Alice A. Han, Giovanni Scardoni, Nate Boggs, Jonathan W. Boyd. An Approach to Investigate Intracellular Protein Network Responses,
Chemical Research in Toxicology, (01 2014): 0. doi: 10.1021/tx400247g

10/01/2013 2.00 Julie A. Vrana, Nathan Boggs, Holly N. Currie, Jonathan Boyd. Amelioration of an undesired action of deguelin,

Toxicon, (11 2013): 0. doi: 10.1016/j.toxicon.2013.07.028

10/01/2013 1.00 Jonathan Boyd, Julie A. Vrana, Holly N. Williams. In vitro approach to predict post-translational phosphorylation response to mixtures,

Toxicology, (11 2012): 0. doi: 10.1016/j.tox.2012.10.010

TOTAL: 3

Number of Pap	pers published in peer-reviewed journals:
	(b) Papers published in non-peer-reviewed journals (N/A for none)
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	(c) Presentations
Number of Pre	sentations: 0.00
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Names of Faculty Supported

NAME	PERCENT_SUPPORTED	National Academy Member
Jonathan Boyd	0.20	
Abel Rodriguez	0.10	
FTE Equivalent:	0.30	
Total Number:	2	

Names of Under Graduate students supported

NAME	PERCENT_SUPPORTED	Discipline
Constance Mitchel	0.01	
David Claypool	0.01	
FTE Equivalent:	0.02	
Total Number:	2	

Student Metrics

This section only applies to graduating undergraduates supported by this agreement in this reporting period

The number of undergraduates funded by this agreement who graduated during this period: 1.00 The number of undergraduates funded by this agreement who graduated during this period with a degree in science, mathematics, engineering, or technology fields:..... 2.00

The number of undergraduates funded by your agreement who graduated during this period and will continue to pursue a graduate or Ph.D. degree in science, mathematics, engineering, or technology fields:..... 2.00

Number of graduating undergraduates who achieved a 3.5 GPA to 4.0 (4.0 max scale):..... 1.00 Number of graduating undergraduates funded by a DoD funded Center of Excellence grant for Education, Research and Engineering:..... 0.00

The number of undergraduates funded by your agreement who graduated during this period and intend to work for the Department of Defense 1.00

The number of undergraduates funded by your agreement who graduated during this period and will receive scholarships or fellowships for further studies in science, mathematics, engineering or technology fields:..... 0.00

Names of Personnel receiving masters degrees

<u>NAME</u>
Total Number:

Names of personnel receiving PHDs

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<u>NAME</u>	
Total Number:	

Names of other research staff

NAME	PERCENT_SUPPORTED	
FTE Equivalent: Total Number:		

Sub Contractors (DD882)

Inventions (DD882)

Scientific Progress

See Attachement

Technology Transfer

Jonathan Boyd, West Virginia University Abel Rodriguez, University of California, Santa Cruz Using Estimations of Entropy to Optimize Complex Human Dynamic Networks under Stress Final Report

Period Covered by the Report June 15, 2012 through December 31, 2013

Date of Report: December 30, 2013

Project Title: Using Estimations of Entropy to Optimize Complex Human

Dynamic Networks under Stress

Contract Number: W911NF-12-1-0165

Total Dollar Value: \$482,835

Program Manager: Dr. Christian Macedonia, Defense Advanced Research Projects

Agency

Submitted by:

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Security Classification – Unclassified Do not mark document as confidential or business proprietary.

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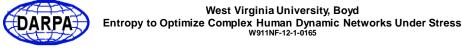
Cc: Dr. Tracy Laabs, <u>tracy.laabs.ctr@darpa.mil</u>, Dr. Kojo Linder, kojo.linder.ctr@darpa.mil, Ms. Jessica Mickey, <u>Jessica.mickey.ctr@darpa.mil</u>.

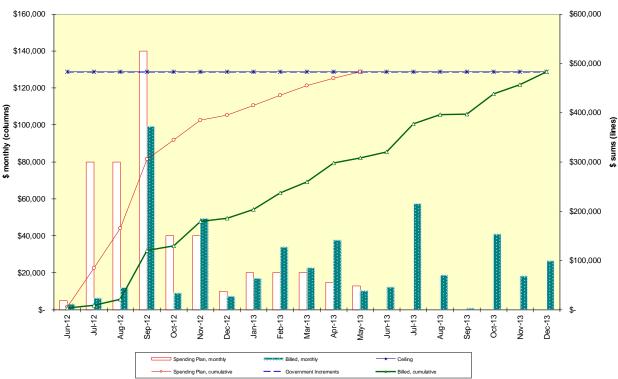
One report to:

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US Army Research Office
Durham, NC 27703

<u>Technical Information – Financial Management</u>

1. Technical Progress / Monthly Expenditure Report (Please provide cumulative spending graph).





Task 1. A written literature report that contains a comparison of biometric sensors that may be beneficial for understanding human response to stress. This will include sensors that are commercially available, academic only, and suggestions for future research. Total cost: \$17,285

Task 2. A written research report that contains preliminary data collection, processing and interpretation of data (from sensors selected in Task 1) that integrates biometric responses of humans under stressful conditions. Further, any computational programs or algorithms used for integration of the data will be included in this report.

Cost to date: \$465,550

Total expenditures for the project - \$482,835

2. Technical Progress / Highlights – Observations

Task 1.1. Literature survey.

This task is complete and was submitted to DARPA/DSO on August 31, 2012.

Task 1.2. Purchase and perform initial T&E; down-select sensors for use in Task 2 This task is complete and was included in our 3rd Quarterly Report.

Task 2.1. Optimize and trouble-shoot data collection for the selected biometric sensors. This task is complete and was included in our 4th Quarterly Report.

Task 2.2. Define the required time to determine baseline.

This task is complete. We have investigated several baselines in order ensure that we have the data to improve the entropy estimations once we start optimizing our entropy calculations. We perform 3 different baselines for EEG:

- 5 minutes high engagement test (picking the correct shape in a sequence)
- 5 minutes low engagement test (clicking spacebar every time that a shape appears, ~ every 2 seconds)
- 5 minutes drowsiness (eyes closes, clicking spacebar every time a chime is heard, ~ every 2 seconds)

and 3 different baselines for all other measurements:

- 5 minutes mental stress (counting backwards from 1000 in intervals of 13)
- 5 minutes relaxation (individual is seated, we typically pull this baseline sample ~ 5 minutes after mental stress activity and ~ 5 minutes before we tell them we are drawing blood)
- 5 minutes of fear/pain stress (before and after venous blood draw)

At this point, our best results have come from data that has been normalized using times taken from 5 minutes mental stress and fear/pain stress. We have explored relaxation, engagement, and drowsiness, but not found any interesting results.

Task 2.3. Experimentally test biometric sensor suite with individuals and teams (with and without stress); optimize calculations of entropy.

EXPERIMENTAL: Currently, testing and analyses of individuals and teams is complete. Data has been collected and cataloged for all biomarkers (see previous Quarterly and Monthly Reports). Additionally, the figures of endogenous and exogenous biomarkers may be found below in Appendix I.

ENTROPY CALCULATIONS: Optimization of entropy model (for individuals and teams) with exogenous biomarkers is complete (see previous Quarterly and Monthly Reports). Additionally, entropy model data is presented in Appendix II below. It should be noted that any of the following calculations (or variants) can be calculated from any exogenous dataset. We chose to explore the following models using heart rate, breathing rate, temperature, activity and posture; additionally, we have performed these calculations with only heart rate, breathing rate and temperature as a means to remove any bias that might be present due to individual activity and posture when performing as a team.

For all subjects, data is downloaded from individual sensors and matched up according to their timestamps in 1 second intervals. We then export the real-time data starting at the first background task (counting backwards from 1000 in intervals of 13) and ending 10 minutes after crime scene house scenario 3 (CSH3) into GraphPad Prism (V5, Cary, NC).

For Shannon Entropy (nee Filter) Slope V1: The exported data from this time interval is then normalized over the time period using a probability distribution. By using probability distribution, the individual's smallest biomarker response (e.g., lowest heart rate, lowest breath rate, etc.) is set as zero and the highest biomarker response (individual's max heart rate, breath rate, etc.) is set to 1; therefore normalizing their individual responses. By including data from the beginning of background data collection through 10 minutes after CSH3 for data normalization, we should have captured their background/baseline exogenous biometric responses, i.e., minimum responses to mental and physical stress (relaxation period) and maximum responses to mental and physical stressors (e.g., count backwards, blood collection, crime scene house scenarios).

For Shannon Norm to Total: The exported data is then normalized over the entire time period for a global maximum value of each exogenous biomarker. We then normalize (divide) each of the exogenous responses (over time) by the respective maximum exogenous biomarker values from the entire experimental time period.

For Shannon Norm to Count: The exported data is then normalized over the entire time period by first isolating a short window of time (up to 180 seconds) during the mental stress activity of counting backwards from baseline exercises. We then normalize (divide) each of the exogenous responses by the respective maximum exogenous biomarker values from this period of mental stress.

For Shannon Norm to Blood: The exported data is then normalized over the entire time period by first isolating a short window of time (up to 180 seconds) during the fear/pain stress activity of having blood drawn during baseline exercises. We then normalize (divide) each of the exogenous responses by the respective maximum exogenous biomarker values from this period of fear/pain stress.

Normalization of the values allows (i.e., all values are on the same scale) for facile comparison between biomarkers directly, which ensures that one biomarker is not weighted more heavily than another.

For all Shannon Variants: After obtaining the normalized responses, we compute the Shannon entropy of each individual exogenous biomarker, $S(\lambda)$, over time (1 second intervals) by taking $S(\lambda) = p(\lambda) \log_2 p(\lambda)$ of each variable, λ .

For Shannon Entropy (nee Filter) Slope V1: Once each variable has its entropy computed over time, we take the Shannon total entropy, S_{Total} , where $S_{Total} = |-\Sigma|S(\lambda)|$, across all variables at each second interval. Additionally, to show Shannon entropy accumulating over time, we take the integral of the sum of Shannon entropy, described as: $S_{Total} = \int S_{Total} dt$.

Finally, to identify discrete changes in total Shannon entropy over time, we take the slope of the total Shannon entropy for a ten second window over time (note: this slope smoothing window may be decreased to a 2 second period if desired without changing the results).

Task 2.4. Write and deliver final report.

The information presented here is intended to serve as the Final Report.

Summary of subjects

	Subject	Gender	Age	Race	Weight (lb.)	Height (in.)	BMI	Body Fat %	Class Year	Rank
	4102	М	22	White	137.6	65	22.9	9.4	Senior	Cadet Captain
Team 1	4103	М	20	White	166.2	74	21.9	12.6	Junior	Cadet 1st Lt.
Tealli 1	4104	М	19	White	207.2	73	27.3	19.6	Freshman	Cadet 4th Class
	4110	М	18	White	155	70	22.2	15.7	Freshman	GMC, 100
	4105	М	19	White	121	67	19	8.2	Sophomore	C/3C
Team 2	4108	М	20	White	196.6	76	23.9	13.3	Sophomore	C/3C
Tealii Z	4112	М	19	White	191.4	75	23.3	17.5	Freshman	C/4C
	4113	М	18	White	200.2	74.5	25.3	21.4	Freshman	C/4C
	4106	М	19	White	152.2	68.5	22.8	14.4	Sophomore	C/3C
Team 3	4111	М	20	White	223.6	76.5	26.9	14.8	Sophomore	C/3C
Tealli 5	4115	М	19	White	183.6	74	23.5	16.6	Freshman	C/4C
	4116	М	19	White	153.8	68.5	23.1	12.5	Sophomore	C/3C
	4107	М	19	White	147.8	68.5	22.2	8.8	Freshman	C/4C
Toam 4	4109	М	19	White	160.6	72.5	21.4	11.7	Freshman	C/4C
Team 4	4114	F	21	White	154.6	66	25	26.5	Junior	C/Colonel
	4117	М	21	White	213.8	72	29	21.6	Junior	C/Lt Col

Relevant Abbreviations

HR = heart rate,

BR = breathing rate,

POS = posture,

ACT = activity,

TEMP K -OR- T = temperature in Kelvin

Ipi = Individual plasma initial

Ipf = Individual plasma final

Tpi = Team plasma initial

Tpf = Team plasma final

Is1 = Individual saliva taken before CSH1 (crime scene house scenario 1)

Is2 = Individual saliva taken after CSH1, but before CSH2 (crime scene house scenario 2)

Is3 = Individual saliva taken after CSH2, but before CSH3 (crime scene house scenario 3)

Is4 = Individual saliva taken after CSH3

Ts1 = Team saliva taken before CSH1

Ts2 = Team saliva taken after CSH1, but before CSH2

Ts3 = Team saliva taken after CSH2, but before CSH3

Ts4 = Team saliva taken after CSH3

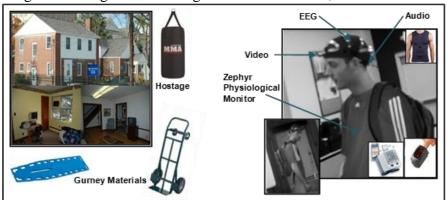
LAY SUMMARY

For all team activities, whether it be sports teams or military units, winning is the ultimate goal. To truly achieve team success, the team should be able to complete a task with the least amount of moves or use the least amount of energy without sacrificing quality.

In this study, we are trying to determine what factors contribute the most to successful teams under stress. We hypothesize that by measuring individual biological responses, such as heart rate, breathing rate, posture, temperature, and saliva/blood components, during various physically and mentally stressful exercises, we will be able to determine the factors that drive overall team success and assemble more effective teams using these factors. Sixteen WVU Air

Force ROTC participants were selected, divided into four different teams of four individuals, their biological responses were monitored (some in real-time and some prior to and immediately following) to a stressful teamwork exercises (mock hostage rescue). Individuals were outfitted with EEG, heart rate, breathing rate, estimated core temperature, sound, activity and posture monitors and tasked with finding the "hostage" and moving it to a safe location; to move the

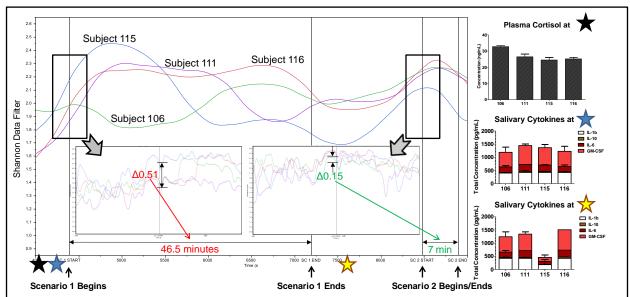
hostage required the assembly of a make-shift gurney from items hidden in the house (see figure at right). These exercises were expected to simulate aspects of the types of stresses endured by U.S. Military Special Operations Forces and Hostage Rescue Teams of



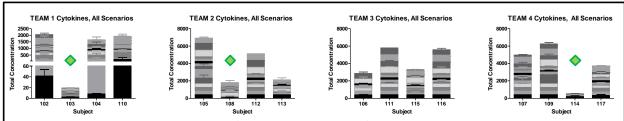
the Federal Bureau of Investigation.

Overall mission recap: During the first mock hostage rescue mission, crime scene house 1 (CSH1), the subjects are told to complete the mission and they will be timed. The CSH1 mission familiarizes the subjects with the crime scene house and the task at hand. For the second hostage rescue mission (CSH2), the subjects are told to go as fast as they can, but still abide by the mission objectives and rules. In the third hostage rescue mission, the subjects are told that they will be timed again, however, 1 minute into the mission an air horn is blown and the mission director states, "Insurgents are returning! You have one minute left or you will be captured!" The second horn blows 1 minute after the first horn and the mission director yells, "FAIL! Assemble the hostage and come back to the garage [rendezvous point]." This mission acts as a failed mission and concludes with a "walk of shame". Blood samples were taken before CSH1 and after CSH3; saliva samples were taken before CSH1, CSH2, CSH3 and after CSH3; blood pressure, temperature, and capillary O2 were also measured before CSH1, CSH2, CSH3 and after CSH3.

As an example, a portion of Team 3 results are shown below. Team 3 was significant because they went from the slowest performance of any team (46.5 minutes; average time for other teams in scenario 1 was 12.5 minutes) to the fastest performance (7 minutes; average time for other teams in scenario 2 was 14.5 minutes). We believe that this team's extreme performance enhancement was due to physiological synchrony of individuals within the team, and with the emergence of a physiological leader (as shown by salivary cytokine concentrations for Team 3 at right) before the second attempt. Note: In post-survey questionnaire, Team 3 did not identify any member as a leader and yet leadership was observed. The concept of salivary cytokine concentrations relating to leadership is supported by additional data from other teams in the second figure below.



Team 3 went from the worst performance to the best performance during a mock hostage rescue scenario based upon time (from 46.5 minutes to 7.5 minutes); Shannon Data Filter (of physiological responses) synchrony increased 3.4x ($\Delta 0.51 \rightarrow \Delta 0.15$). The non-smoothed Shannon Data Filter responses are shown in the small inset boxes; note: synchrony is achieved by all subjects increasing their response to match Subject 115. Salivary concentrations of cortisol and cytokines (IL-1b, IL-10, IL-8, GM-CSF) are shown at right for each subject; note: prior to Scenario 2, Subject 115 showed ~3X decrease in salivary cytokines when compared to other subjects in his team.



Aggregated salivary cytokine concentrations (summed for all scenarios) for 4 different teams. In post-scenario surveys, subjects labeled with \diamondsuit were identified as primary leaders of the team; team 3 indicated that there was no leader. Lower salivary cytokine responses appear to be related to leadership within a team.

LINKING BIOMARKERS TO MISSION SUCCESS

The mock hostage rescue missions that were performed by each subject as an individual and as part of a four person team at the WVU crime scene complex provided some extremely useful information about subject performance, leadership, and task efficiency. To understand how these metrics correspond to physiologically relevant biomarkers in blood and saliva, correlation analysis was performed in SAS jmp. Statistically significant correlations between three different metrics (performance, leadership, and efficiency) and subject biomarkers are shown in Table 1 below. The pairwise correlation analysis compared each pair of variables across all subjects (N=16). For some variables, N=15 because one subject did not provide enough saliva sample during the team tasks to perform all assays necessary. Additionally, during the individual scenarios, one subject did not have their body fat % recorded, therefore N=15 for individual correlations involving body fat %. Any correlations with P > .05 were discarded. Due to potential statistical bias (because all individuals on a team finished at the same time), team performance was not calculated for pairwise correlation analysis. In Table 1, individual plasma initial and individual plasma final are abbreviated as ipi and ipf, respectively. Team plasma initial and team plasma final follow the same style, tpi and tpf, respectively. For GSH, abbreviations are "individual initial (ii), individual final (if), team initial (ti), and team final (tf)," because GSH was measured in the erythrocyte lysate (i.e. intracellular components of red blood cells). For salivary samples, the abbreviation style is the same, where individual saliva sample 1 is "is1" and team saliva sample 1 is "ts1," and so forth.

The sampling collection times for individual and team missions follow the scheme below: Plasma/blood Initial --> Saliva 1 --> CSH1 --> Saliva 2 --> CSH2 --> Saliva 3 --> CSH3 --> Saliva 4--> Plasma/blood Final

The variables highlighted in RED were negatively correlated, while those in GREEN represent positively correlated variables.

Table 1.

Variable		PERFORM	ANCE				
CSH2 time (indiv)	Variable	_		N	Lower 95%	Upper 95%	P value
CSH2 time (indiv)		ř					
SSH2 time (indis)					1		
CSH2 ime (indity)							
CSH1_time (indiv)	CSH3_time (indiv)	BMI	-0.4978	16	-0.7969	-0.0027	0.0498
CSEIL_time (indiv)	CSH2_time (indiv)	Body_fat_%	-0.626	15	-0.8619	-0.1675	0.0125
Indiv. Containine	CSH3_time (indiv)	Body_fat_%	-0.5907	15	-0.8467	-0.1125	0.0204
CSH3_time (indiv)	_ ` ′	Chloride_ipi		16	1		
Variable	_			_	1		
Variable	CSH3_time (indiv)			16	0.0173	0.8021	0.0442
Leadership Score CSH1		LEADERS			,		
Leadership Score CSH1							
Leadership Score CSH1							
Leadership Score CSH CSH Los Los		- 1					
Leadership Score_Overall CSH_ti -0.4992 16 -0.7999 -0.0111 0.045 Leadership Score_Overall IPNg_183 -0.59314 16 -0.813 -0.0485 0.0341 Leadership Score_Overall IPNg_183 -0.5838 16 -0.8872 -0.124 0.017 Leadership Score_CSH1 II.1b_181 -0.6571 16 -0.8696 -0.2394 0.0057 Leadership Score_CSH1 II.1b_182 -0.5885 16 -0.8802 -0.1013 0.0216 Leadership Score_CSH1 II.1b_182 -0.5885 16 -0.8802 -0.1013 0.0216 Leadership Score_CSH1 II.1b_183 -0.6217 16 -0.8842 0.1812 0.0101 Leadership Score_CSH2 II.1b_183 -0.5119 16 -0.8037 -0.018 0.0426 Leadership Score_CSH2 II.1b_183 -0.5119 16 -0.8037 -0.0218 0.0426 Leadership Score_CSH1 II.6_181 -0.6655 16 -0.8693 -0.2384 0.0037 Leadership Score_CSH1 II.6_182 -0.5728 16 -0.8322 -0.1077 0.0204 Leadership Score_CSH1 II.6_182 -0.5728 16 -0.8322 -0.1077 0.0204 Leadership Score_CSH1 II.6_183 -0.5269 16 -0.8099 -0.7044 0.0114 Leadership Score_CSH1 II.6_183 -0.5206 16 -0.8078 -0.0335 0.0387 Leadership Score_CSH1 II.6_183 -0.5206 16 -0.8078 -0.0335 0.0387 Leadership Score_CSH1 II.8_183 -0.5629 16 -0.8078 -0.0335 0.0387 Leadership Score_CSH1 II.8_184 -0.533 16 -0.8123 -0.0466 0.0347 Leadership Score_CSH1 II.8_184 -0.533 16 -0.8123 -0.0466 0.0347 Leadership Score_CSH2 Testosterone_Ipi -0.5187 16 -0.8029 -0.0309 0.0395 Wariable Development of the state				_			
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Leadership Score_CSH1	. –		-0.5283	16	-0.8115	-0.0442	0.0354
Leadership Score_Overall II.1b_ts3		IL1b_ts3	-0.6217	16	-0.8542	-0.1821	0.0101
Leadership Score_CSH1	Leadership Score_CSH2	IL1b_ts3	-0.5119	16	-0.8037	-0.0218	0.0426
Leadership Score_Overall	Leadership Score_overall	IL1b_ts3	-0.6565	16	-0.8693	-0.2384	0.0057
Leadership Score_overall IL6_ts2		IL6_ts1		16			0.0114
Leadership Score CSH1				16			
Leadership Score_overall IL6_ts3							
Leadership Score CSH1				-			
Leadership Score_CSH1				-			
Description Control							
Variable		_		-			
Variable by Variable Correlation N Lower 95% P value Activity/time(s)_CSH1_indiv ACTH_ipf -0.5083 16 -0.802 -0.0169 0.0444 Activity/time(s)_CSH1_indiv ACTH_ipi -0.5576 16 -0.8252 -0.0856 0.0248 Activity/time(s)_CSH2_indiv ACTH_ipi -0.5838 16 -0.8372 -0.124 0.0176 Activity/time(s)_CSH1_indiv Cl_ipi -0.7502 16 -0.9082 -0.4052 0.0008 Activity/time(s)_CSH1_indiv Cl_ipi -0.6728 16 -0.9763 -0.2657 0.0043 Activity/time(s)_SumCSH123_indiv Cl_ipi -0.7573 16 -0.904 -0.0226 0.0423 Efficiency (team) Variable by Variable Correlation N Lower 95% P value Activity/time(s)_CSH1_team Cortisol_tpf -0.5012 16 -0.7985 -0.0073 0.048 Activity/time(s)_CSH1_team GM-CSF_ts1 -0.6473 16 -0.822	Leadership Beore_eBriz	<u> </u>		10	-0.0007	-0.0507	0.0373
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$\frac{1}{1}$	Activity/time(s)_CSH1_team	TNFa_ts1	-0.547	16	-0.8203	-0.0703	0.0283
Activity/time(s)_CSH2_team TNFa_ts2 0.5553 16 0.0823 0.8242 0.0255	Activity/time(s)_CSH2_team	TNFa_ts2	0.5553	16	0.0823	0.8242	0.0255

PERFORMANCE: For the mock hostage rescue missions, the length of time to complete each mission (CSH1, CSH2, or CSH3) was used to describe subject performance (i.e. advanced performance would be the shortest amount of time to complete the mission and poor performance would be the longest amount of time to complete the mission). In CSH1, the subjects are familiarizing themselves with their surroundings in the context of the tasks. Initial chloride concentrations were positively correlated with CSH1 time (i.e. low concentrations of chloride were correlated with shorter amount of time to complete CSH1), showing that initial chloride levels may be predictive of CSH1 performance. The best mission/scenario to determine subject performance would be CSH2, where the subjects are informed to go as fast as possible. For this mission/scenario, ACTH was positively correlated with CSH2 performance, whereas BMI and body fat % were negatively correlated to mission performance . BMI and body fat % negatively correlated to CSH2 performance is a surprising finding- the higher your BMI or body fat %, the faster you went (took less time to complete CSH2). Performance during the failure task (CSH3) is positively correlated to salivary cortisol concentrations, whereas BMI and body fat % were negatively correlated to CSH3 performance (similar to CSH2).

LEADERSHIP: While completing the mock hostage rescue scenarios/missions as part of a four person team, subjects were given a simple post-mission survey to be completed after each mission (CSH1, 2, and 3) and an "overall" survey to describe their own assessment of performance, communication, leadership, etc. In this survey, subjects were asked to state who on their team was the leader(s) for each mission, and who was the overall leader (consistent leader for all missions). To score these responses, the subject that was indicated

Leadership Scores								
Team & Subject	CSH1	CSH2	CSH3	Overall Leadership				
Team 1								
102	1.5	0.5	0.5	0.5				
103	2.5	0.5	0.5	1.5				
104	0	0	0	0				
110	0	0	0	0				
Team 2								
105	0	0	0	0				
108	3	3	4	4				
112	0	0	0	0				
113	0	0	0	0				
Team 3*								
106	0	0	0	0				
111	0	0	0	0				
115	0	0	1	0				
116	1	1	0	1				
Team 4								
107	0	0	0	0				
109	0	0.5	0.5	0				
114	1.5	2	0.5	3				
117	0.5	0.5	0	0				

as the leader received a score of 1, totaling 4 if all subjects selected the same individual as their leader. Some subjects indicated 2 people as the leader for a particular mission, therefore their score was recorded as 0.5 (i.e. 0.5 for subject X and 0.5 for subject Y if a team member said "subjects X and Y were leaders for this mission"). The "Overall leader" leadership score is not a

sum of the three missions; rather it is an independent score from the overall mission success survey where subjects answered the question, "Was there an apparent leader of the group overall [encompassing all 3 missions]?" The asterisk on Team 3 corresponds to their leadership scores indicating no real leader; however from post-video analysis, we can see that subject 111 was more of a social leader and subject 115 appeared to be a physiological leader. From these leadership scores (shown above), we performed correlation analysis on the leadership scores with blood/plasma and salivary biomarkers. In Table 1, initial GSH blood concentrations are inversely proportional to CSH1 and overall leadership scores (lower blood GSH concentrations are related to higher leadership scores), implicating GSH as a predictive biomarker of demonstrated leadership. Also, final ACTH plasma concentrations are inversely proportional to CSH2 leadership scores (the task where the teams are told to go as fast as they can). Finally, various salivary markers, such as GM-CSF, IFN-γ, IL1b, IL6, IL8 as well as plasma testosterone are all inversely proportional to CSH1, CSH2, and/or overall leadership scores; thus subjects with a high leadership score have lower salivary or plasma concentrations of these biomarkers. However, for the "failed" mission (CSH3), leadership scores were proportional to final plasma glucose concentrations (taken immediately following completion of CSH3).

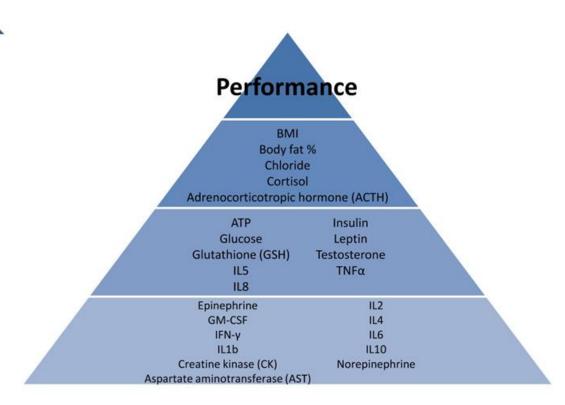
EFFICIENCY: The zephyr external monitor measures several metrics, one of which is activity. To determine subject efficiency during a given mission, we summed an individual's activity during that scenario and divided by the time it took to complete that scenario/mission (i.e. total activity for CSH1 divided by total time to complete CSH1). Since we described activity/time in this manner, efficiency can also be called average activity per second. A high activity/time value could represent low efficiency (inefficiency), where many movements were needed to complete the mission. These calculations were performed for missions where subjects completed the scenarios/missions as an individual (Efficiency individual) and team efficiency was calculated by summing the individual's activity for each scenario and dividing by their respective team's time to complete a given task (i.e. subject 102's summed activity for CSH1(team) was divided by the time it took for team 1 to complete CSH1, since 102 is on team 1). As an individual, efficiency was correlated with subject initial and final ACTH plasma concentrations, initial chloride plasma concentrations and cortisol 3 (collected after CSH2) salivary concentrations. However, these biomarkers were not significantly correlated to efficiency when the subjects were part of a four person team; this may be due to the distribution of work when part of a team and varied leadership roles. As a team, efficiency was correlated with subject initial and final plasma cortisol concentrations and several salivary cytokines (GM-CSF, IFN-γ, IL1b, IL4, IL6, IL8, TNF α) as well as initial saliva testosterone concentrations.

Determining biomarker signaling hierarchy via pairwise correlation analysis

In order to determine the biological significance and a hierarchy of biological importance to mission-success parameters, such as performance, leadership, and efficiency, we performed pairwise correlation analysis across all biomarkers measured, and structured them as they are related to the mission-success parameters of interest. Below, we have described the biomarker signaling hierarchy in relation to mission performance, leadership, and mission efficiency. The biomarker signaling hierarchy can be visualized with a pyramid, where the first tier is correlated to the second tier, the second tier is correlated to the third tier, and the third tier is correlated to

the fourth tier. Specific correlation connections can be found in the attached excel file "correlation indiv_performance".

Biomarker signaling hierarchy







Efficiency (individual)

Chloride Cortisol Adrenocorticotropic hormone (ACTH)

 ATP
 IL8

 BMI
 Insulin

 Body Fat %
 Testosterone

 IL5
 TNFα

Aspartate aminotransferase (AST) Glutathione (GSH) IL6
Creatine Kinase (CK) IFN-γ IL10
Epinephrine IL1b Leptin
Glucose IL2 Norepinephrine
GM-CSF IL4

Efficiency (team)

GM-CSF IFN-y
Cortisol IL1b
IL4 IL6
Testosterone IL8
TNF α

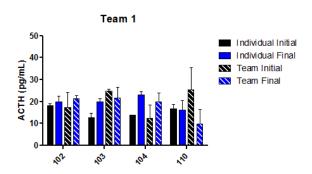
ATP IL2
Creatine kinase (CK) IL5
Chloride IL10
Epinephrine Norepinephrine

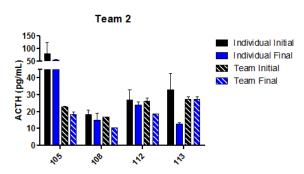
BMI Glucose
Body Fat % Glutathione (GSH)
Leptin Adrenocorticotropin hormone (ACTH)

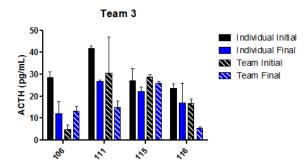
As you can see from efficiency (team) and leadership hierarchy pyramids, the biometrics contributing to these desired endpoints are very similar. This means that some of these biomarkers are driving both leadership and individual efficiency while on a four person team. However, it is important to note that leadership scores and team efficiency is not correlated to each other. Again, this shows that while characteristics/biomarkers indicative of leadership are also related to efficiency, leadership itself (as tabulated from post-scenarios surveys) is not related to efficiency. Could some combination of leadership biomarkers and efficiency biomarkers be indicative of superior mission performance? Further experimentation and model development are necessary to combine these metrics into overall advanced mission success.

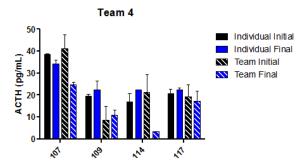
APPENDIX I: ENDOGENOUS BIOMARKER DATA

Individual and Team Plasma ACTH

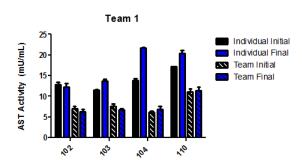


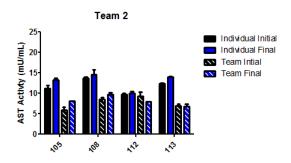


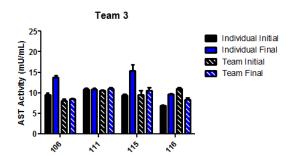


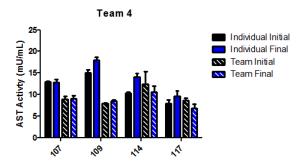


Individual and Team Plasma AST

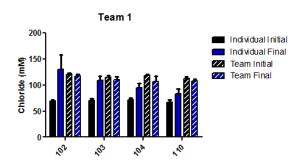


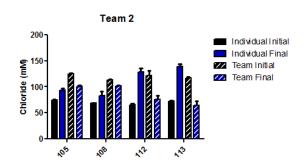


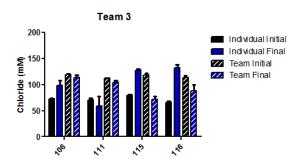


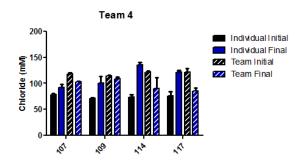


Individual and Team Plasma Chloride

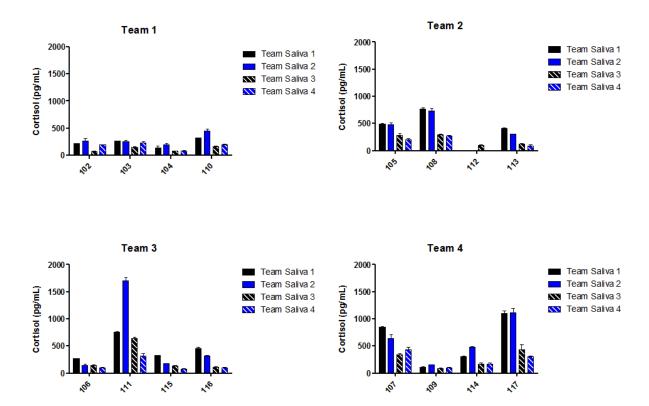




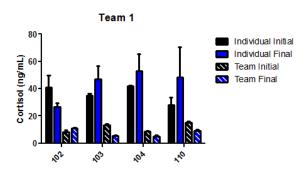


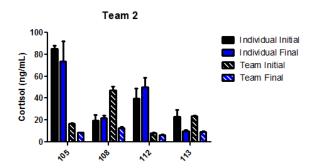


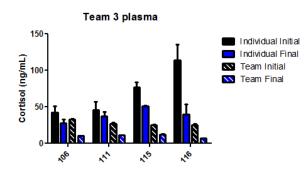
Team Saliva Cortisol

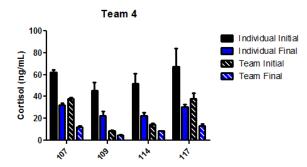


Individual and Team Plasma Cortisol

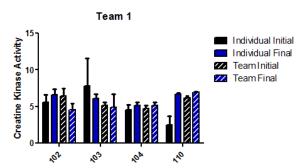


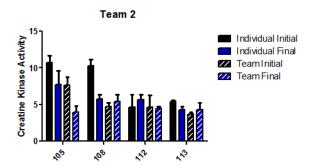


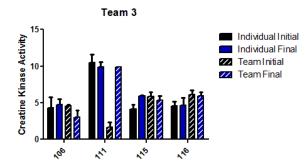


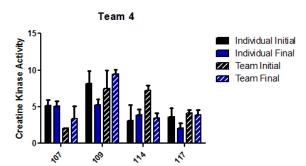


Individual and Team Plasma Creatine Kinase

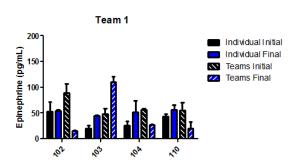


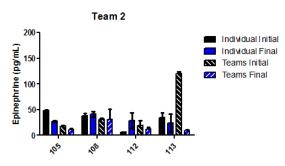


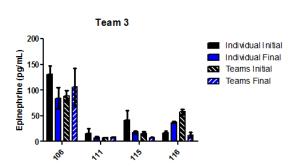


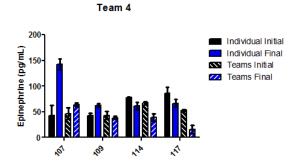


Individual and Team Plasma Epinephrine

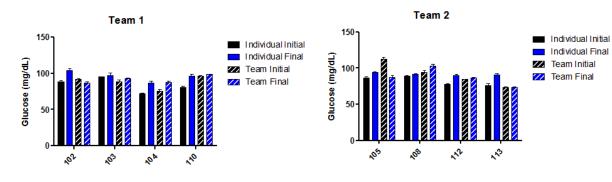


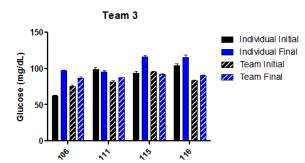


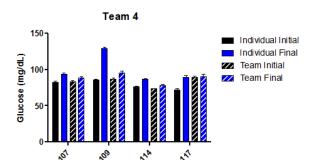




Individual and Team Plasma Glucose

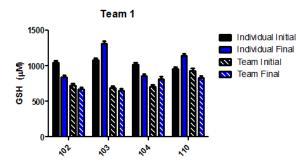


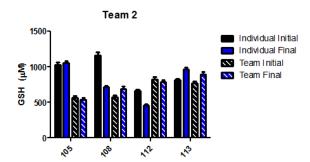


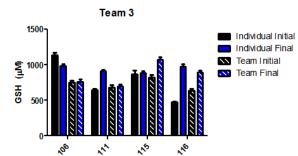


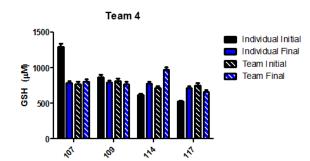
Team Final

Individual and Team Plasma GSH

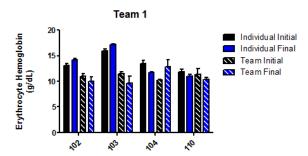


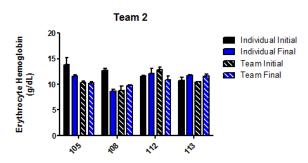


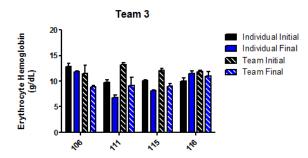


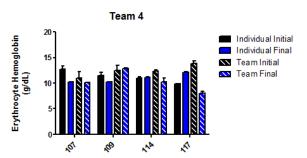


Individual and Team Erythrocyte Hemoglobin

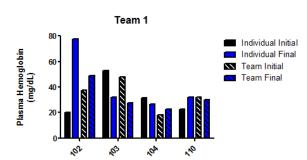


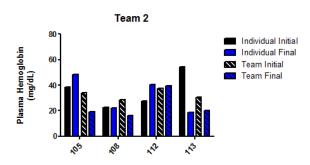


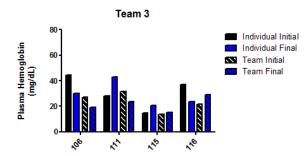


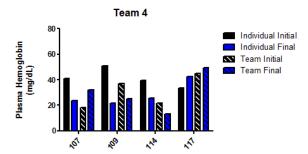


Individual and Team Plasma Hemoglobin

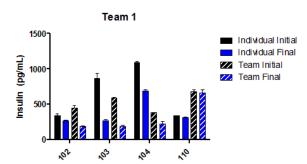


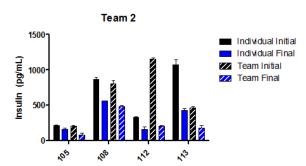


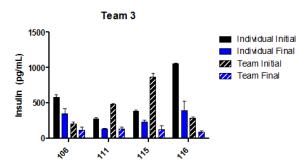


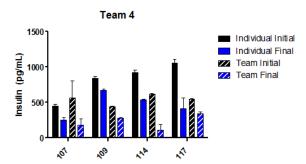


Individual and Team Plasma Insulin

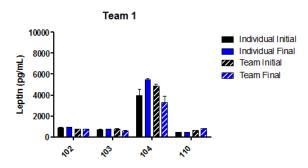


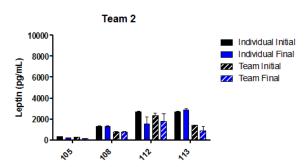


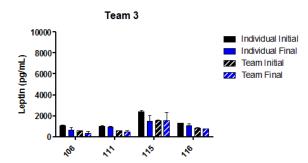


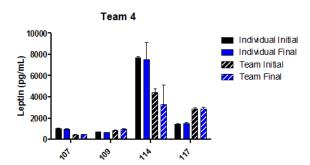


Individual and Team Plasma Leptin

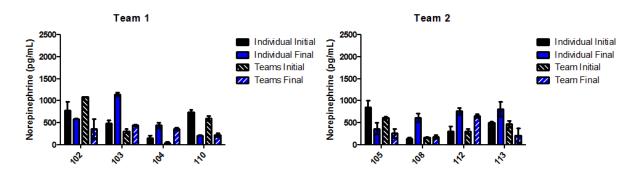


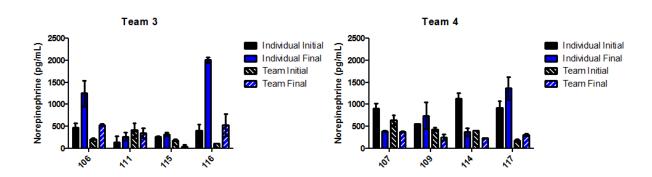




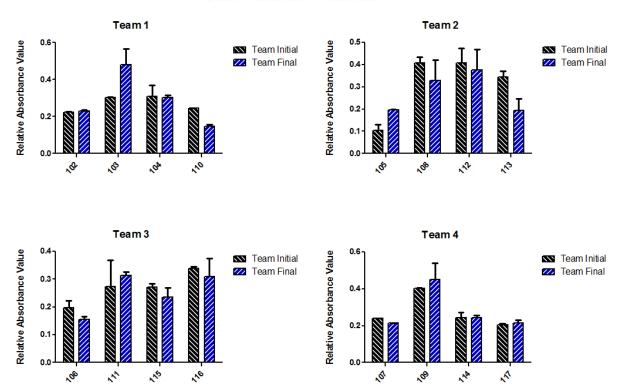


Individual and Team Plasma Norepinephrine

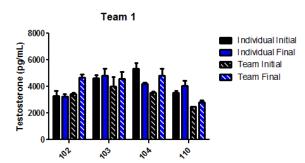


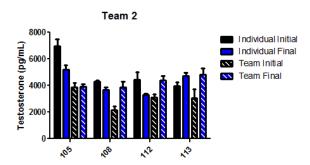


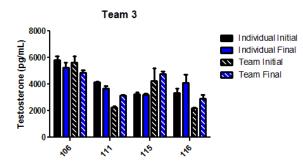
Team Plasma Prolactin

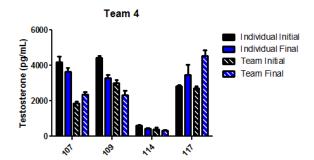


Individual and Team Plasma Testosterone

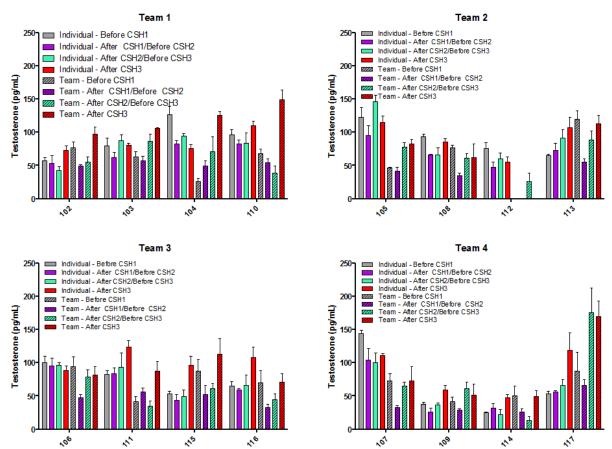


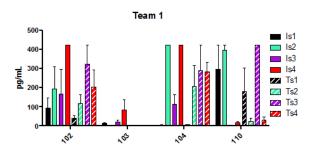


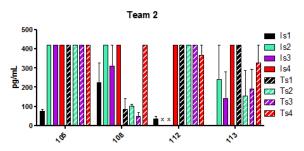


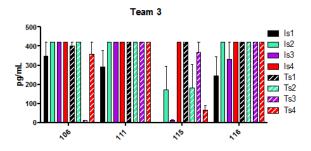


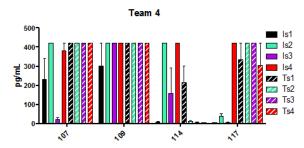
Individual and Team Saliva Testosterone

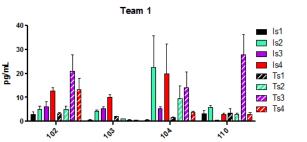


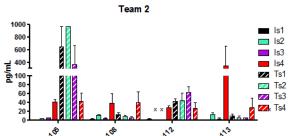


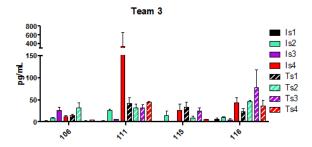


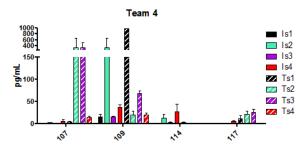


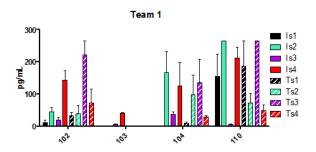


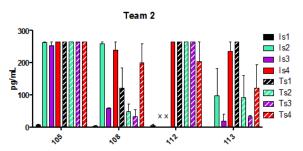


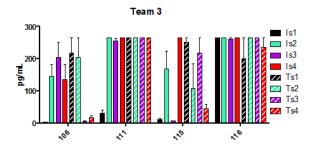


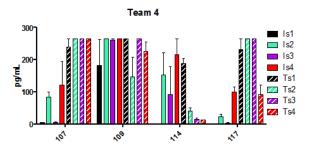


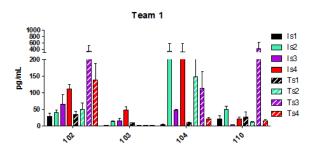


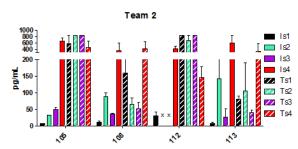


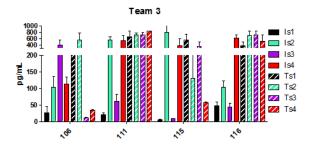


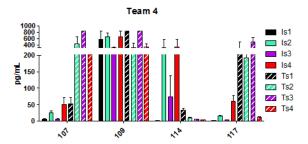


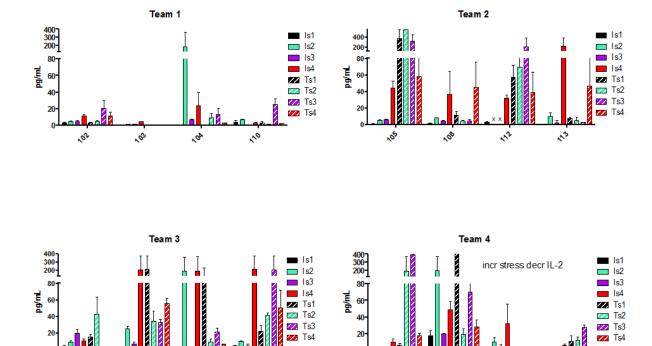


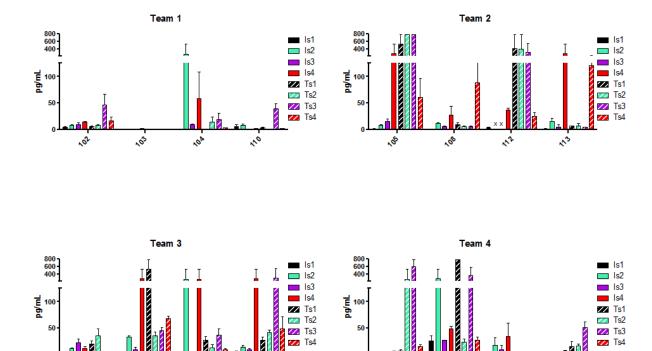


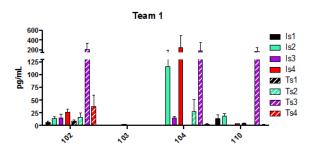


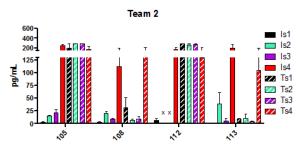


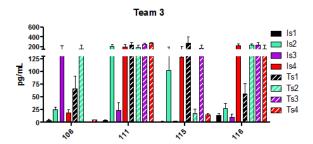


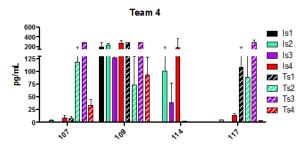


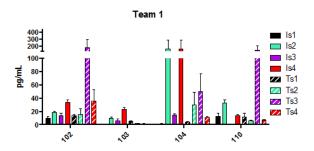


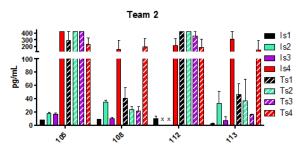


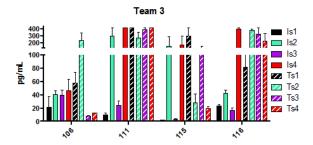


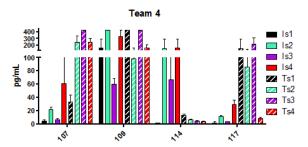




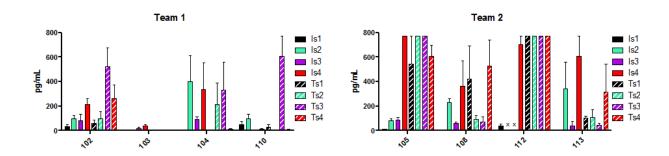


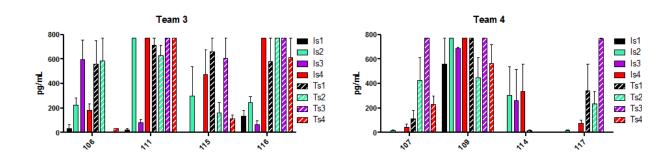




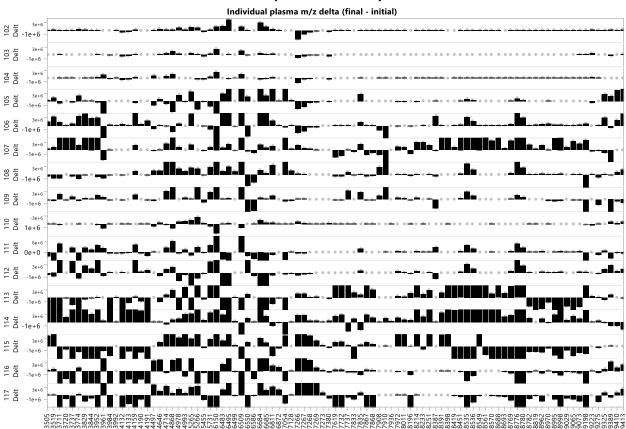


Individual and Team Saliva GM-CSF





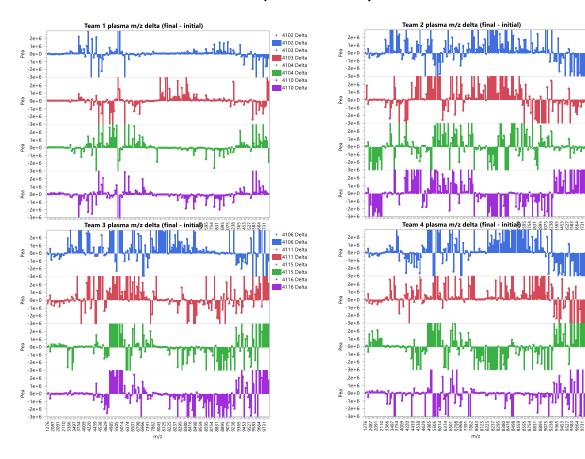
Individual Mass Spectrometry Metabolome



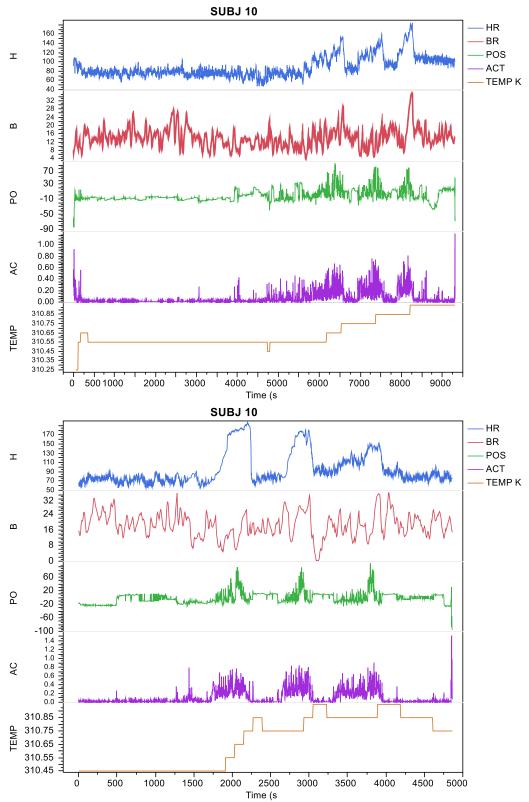
Team Mass Spectrometry Metabolome

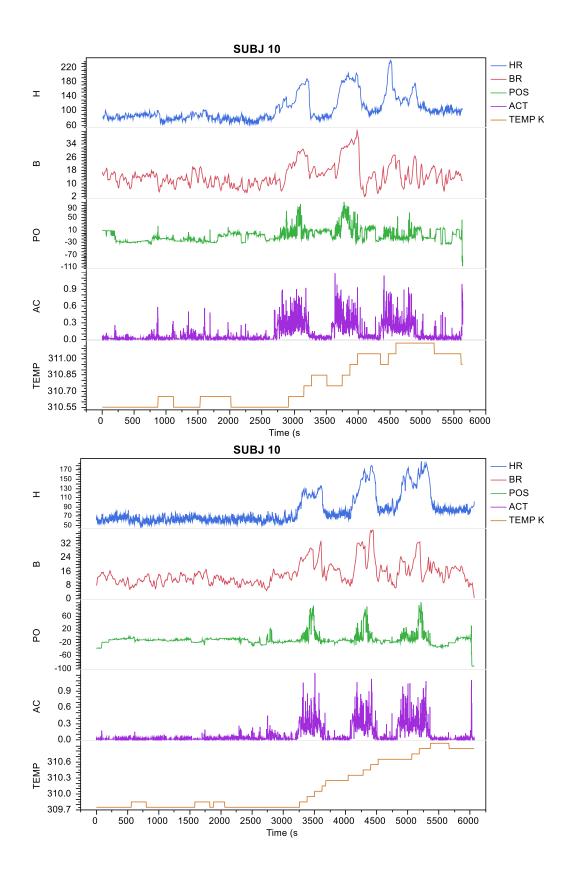
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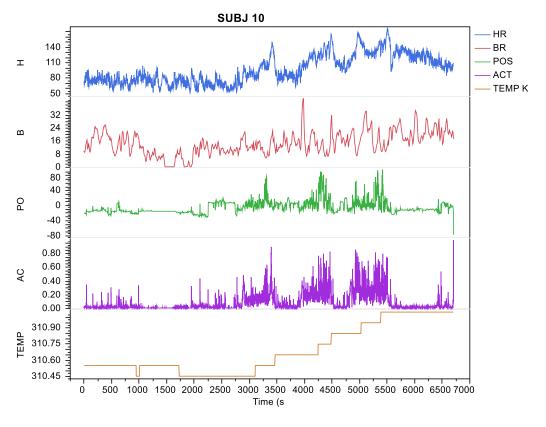
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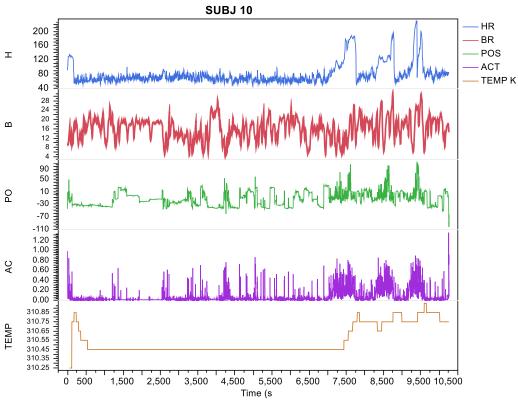


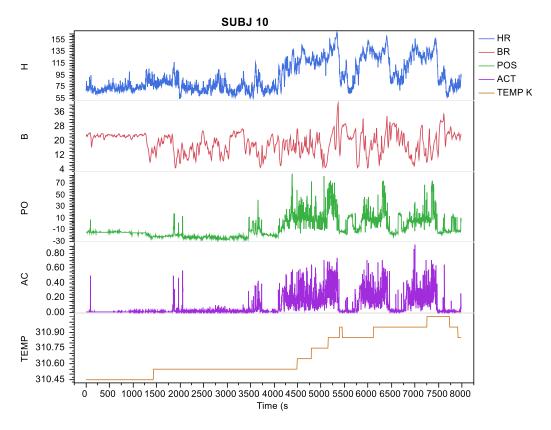
EXOGENOUS BIOMARKER DATA FROM INDIVIDUAL EXPERIMENTS

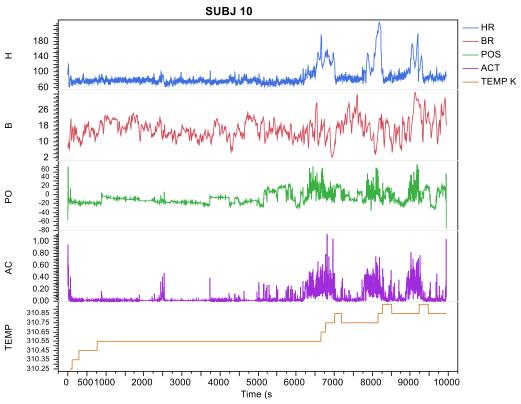


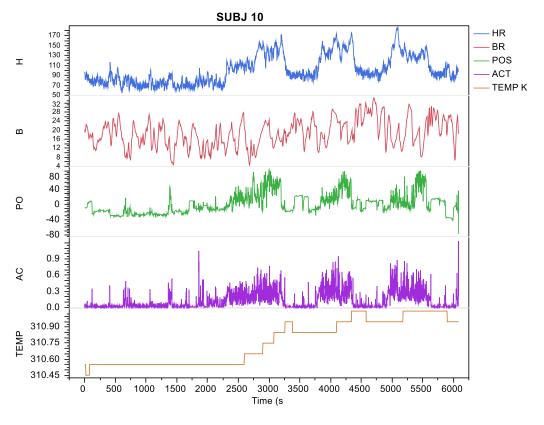


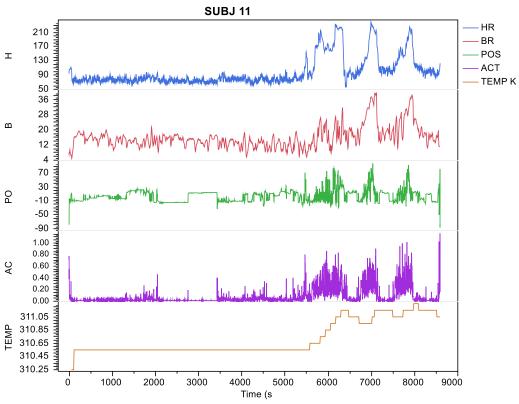


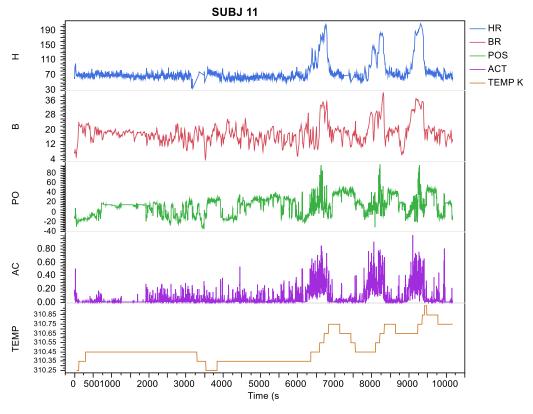


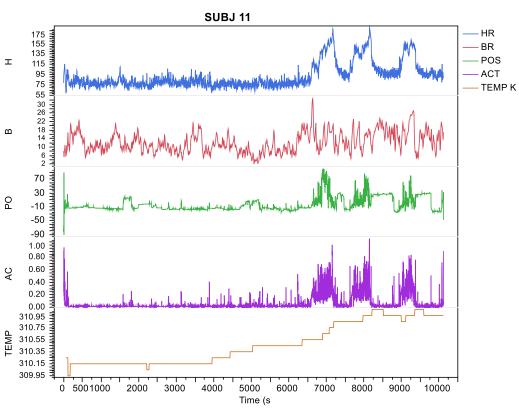


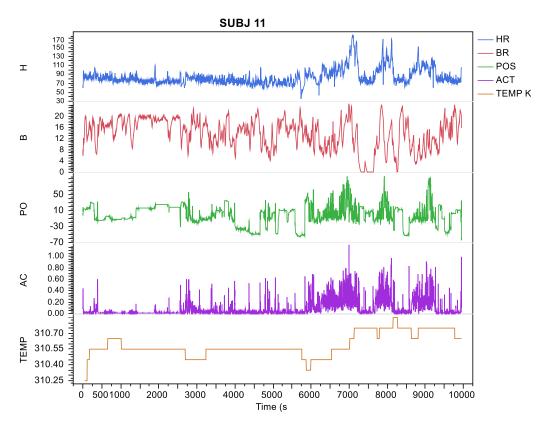


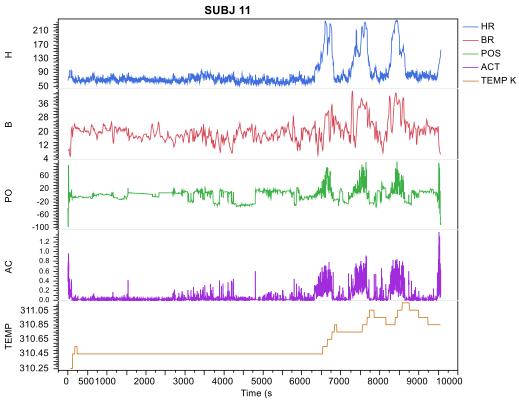


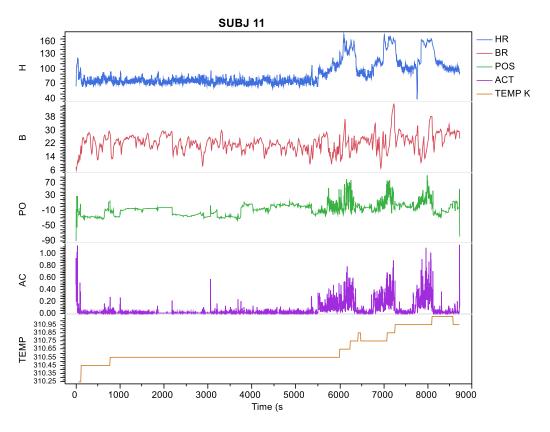


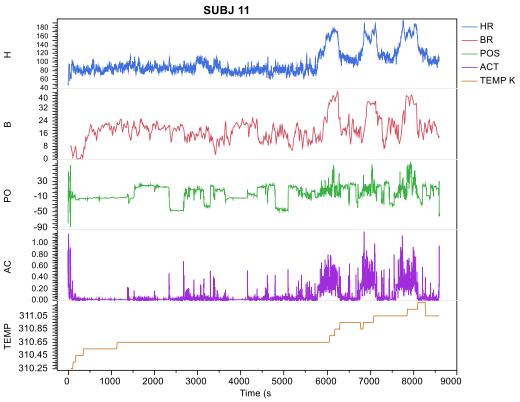


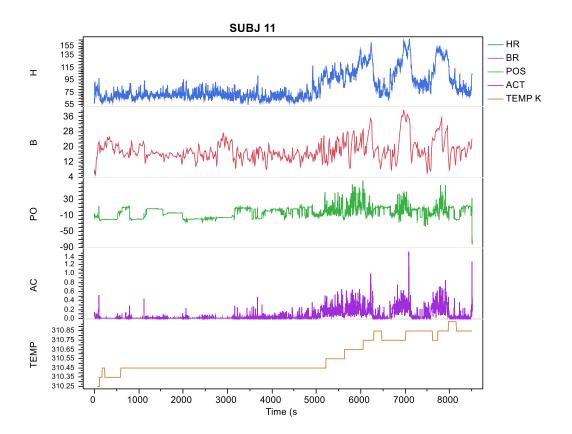








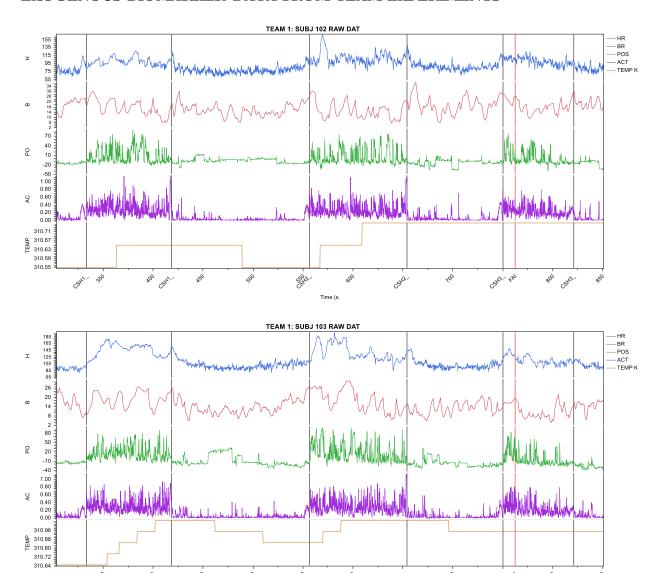




EXOGENOUS BIOMARKER DATA FROM TEAM EXPERIMENTS

CSH1/

400



E.

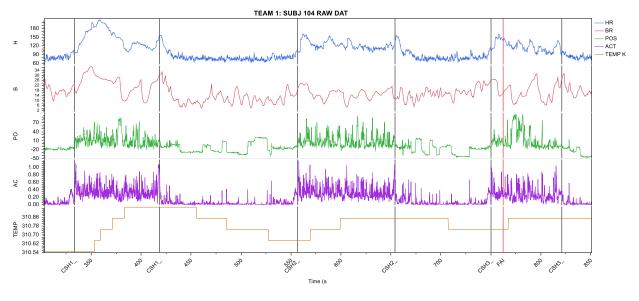
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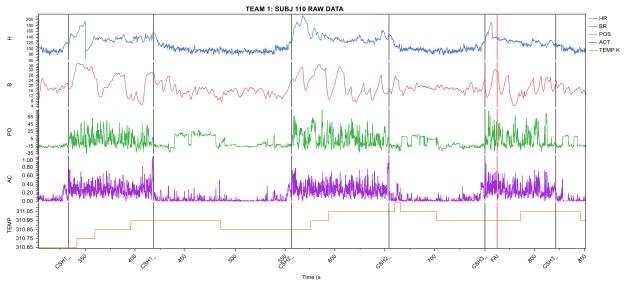
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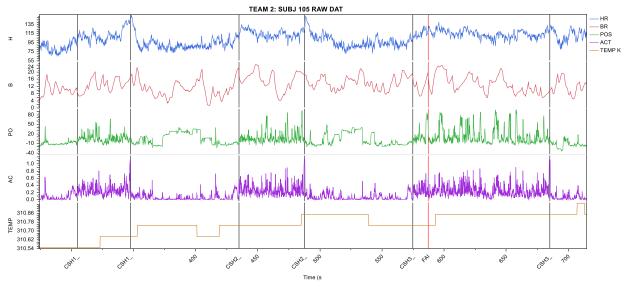
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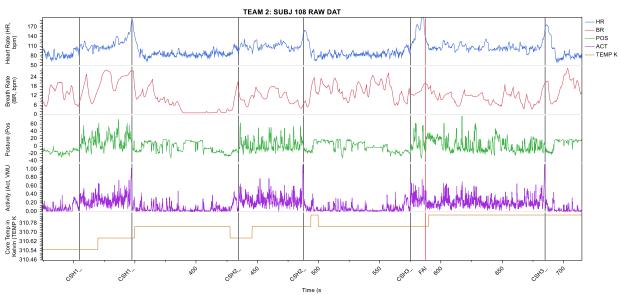
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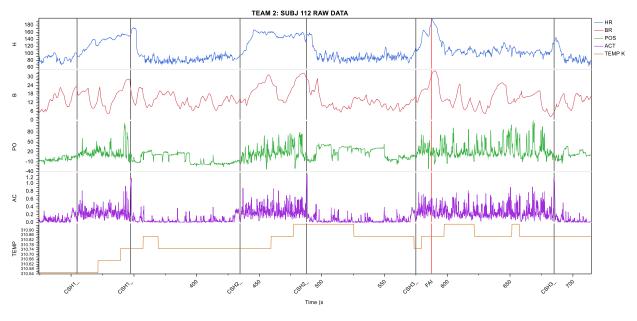
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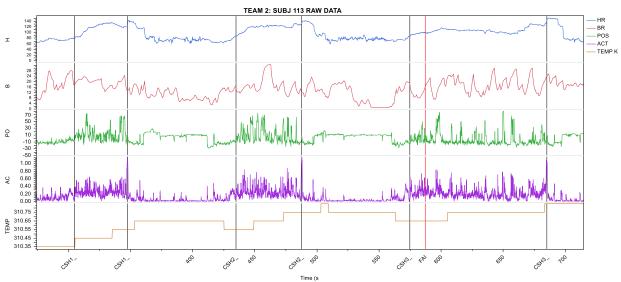


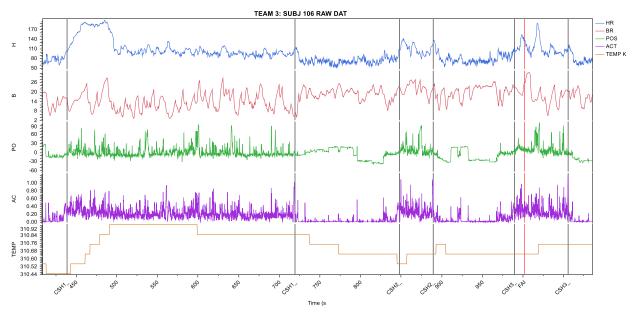


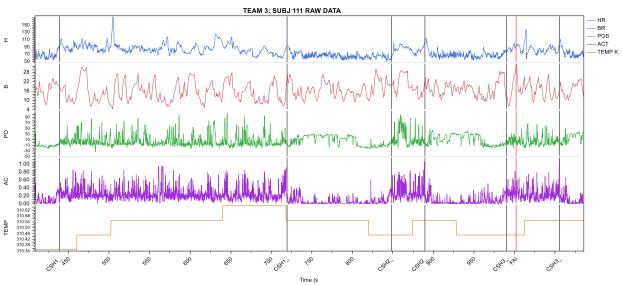


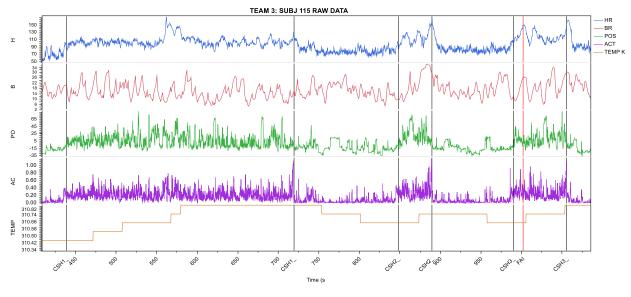


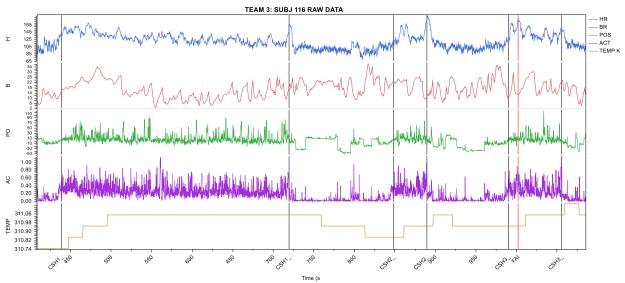


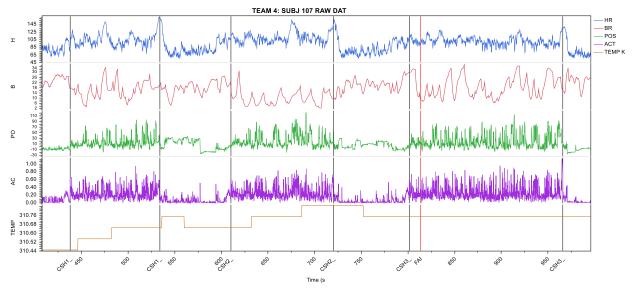


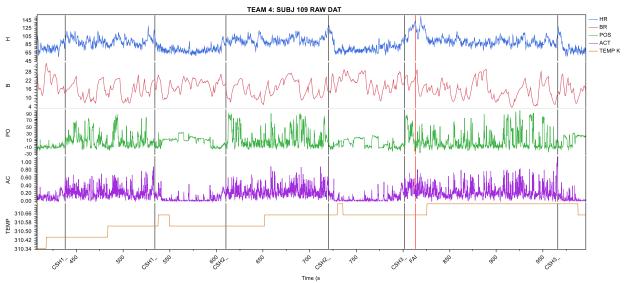


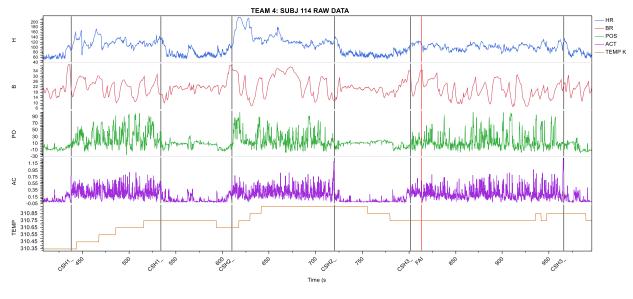


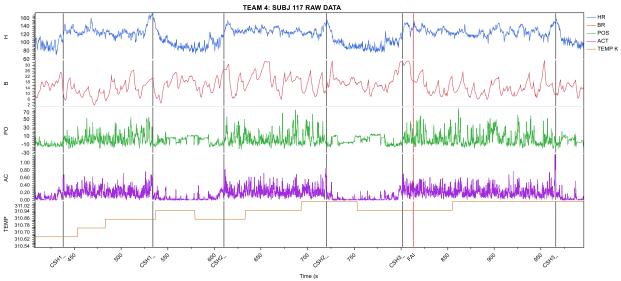




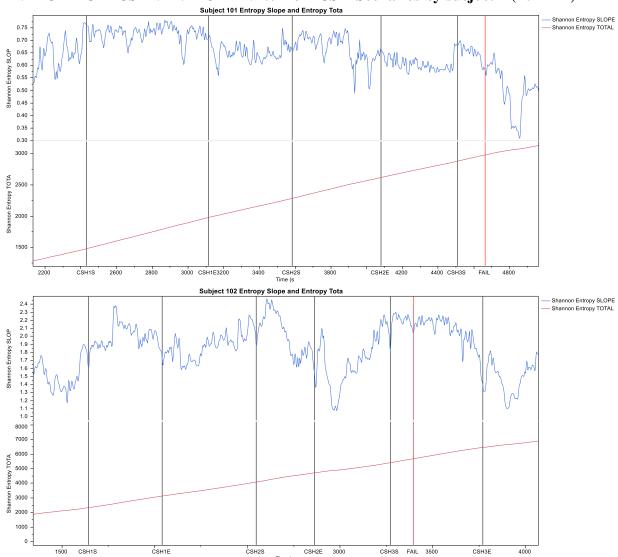


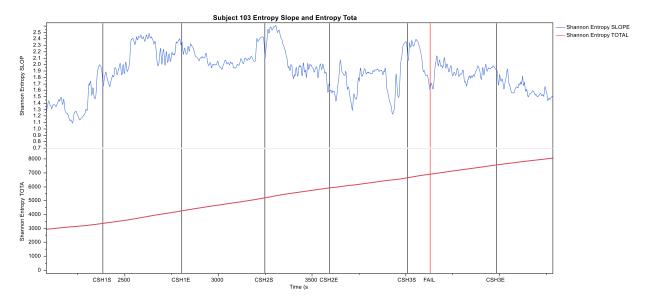


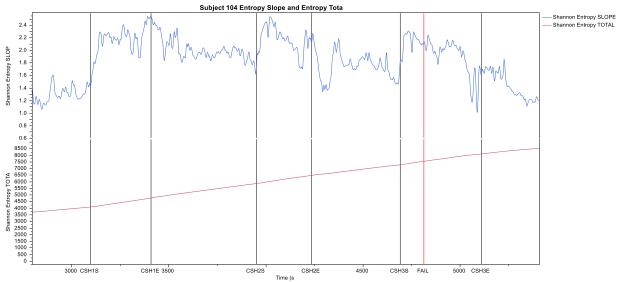


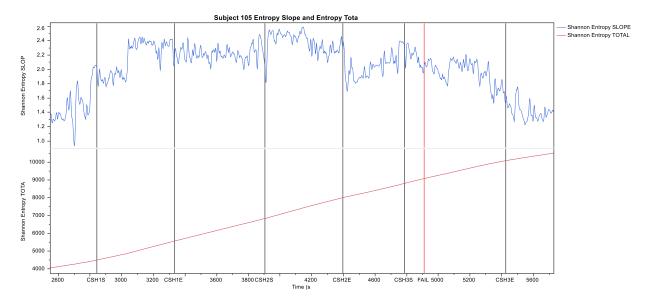


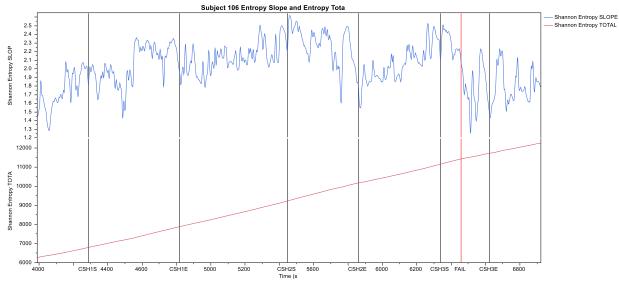
APPENDIX II: ENTROPY CALCS INDIVIDUAL V1.0 from CSH Scenarios by subject # (101-117)

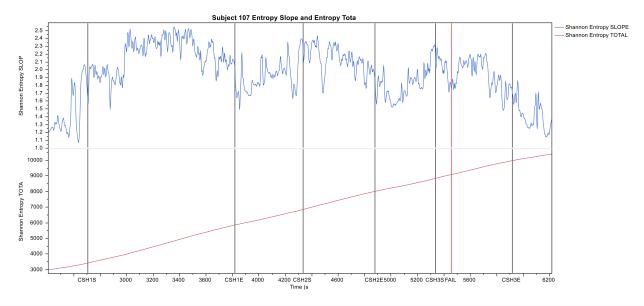


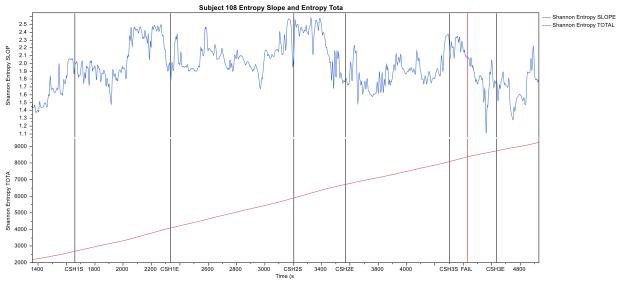


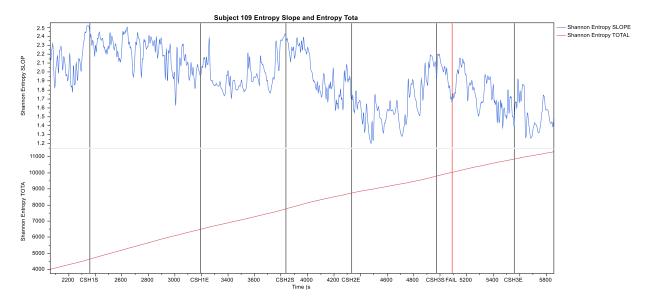


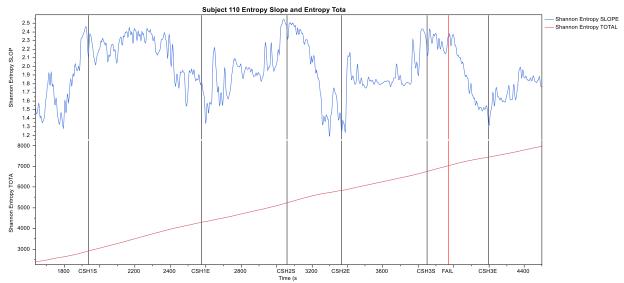


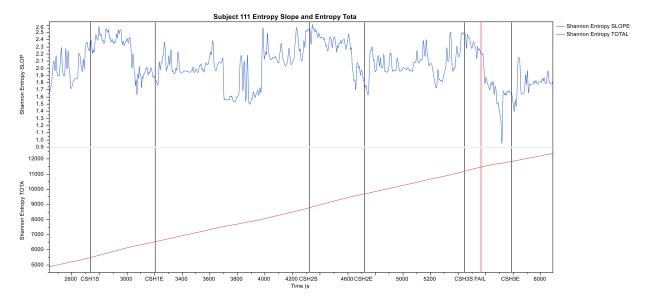


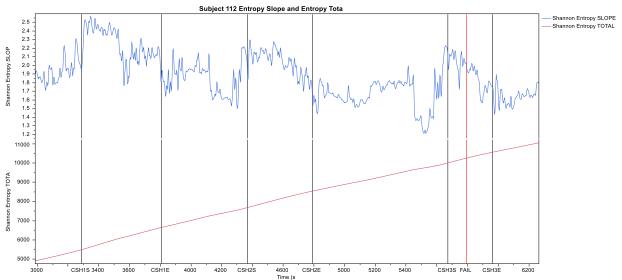


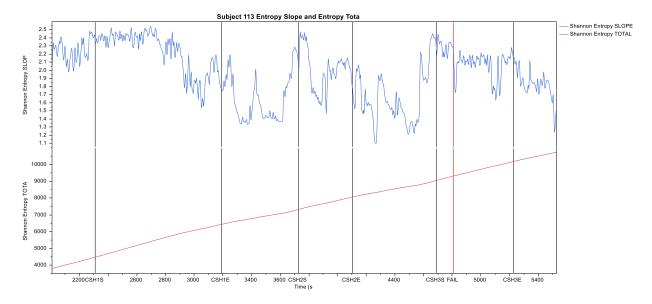


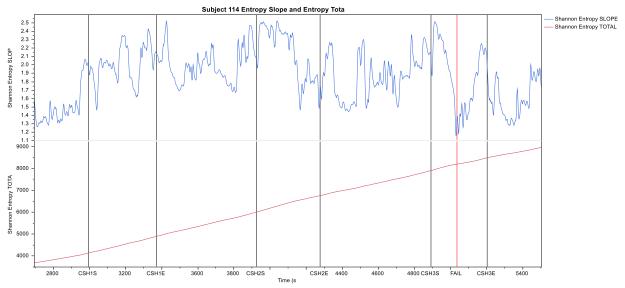


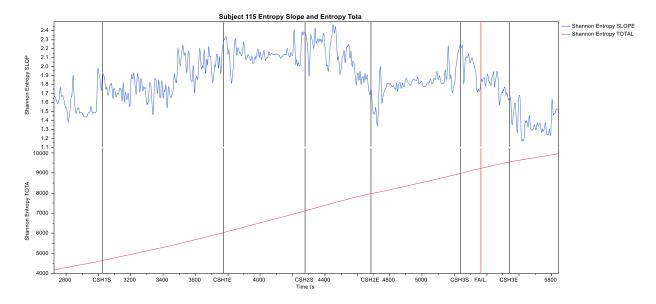


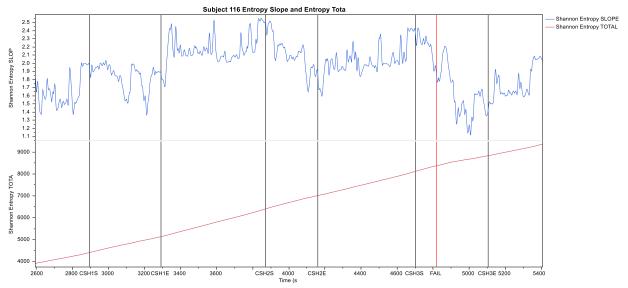


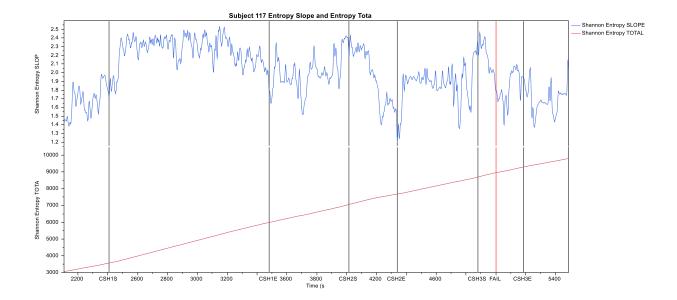




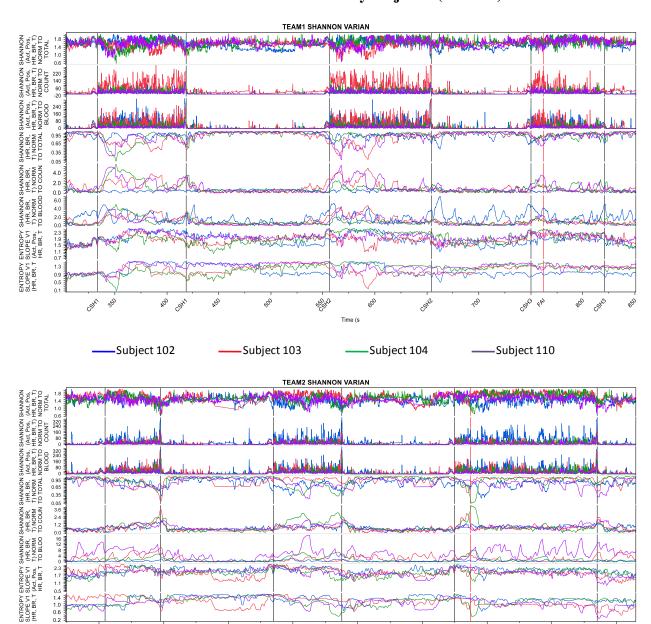




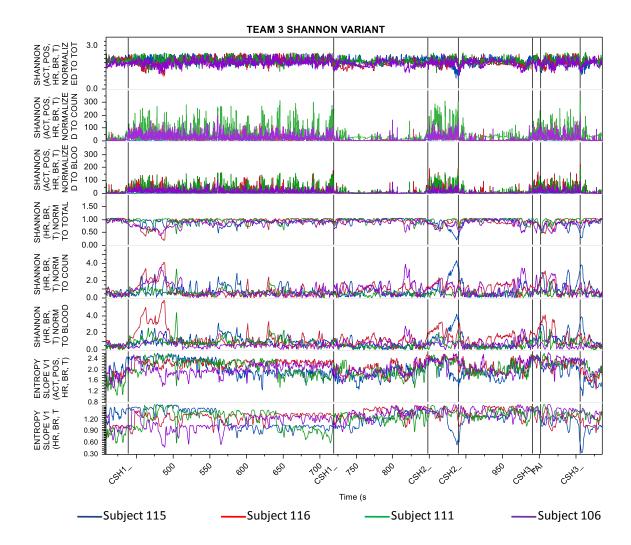


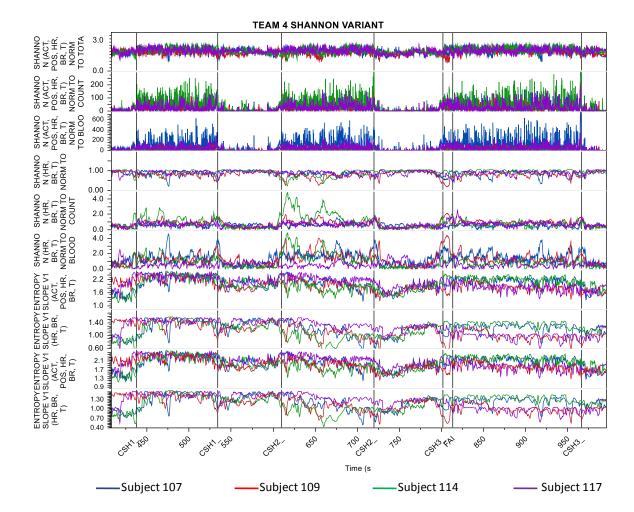


ENTROPY CALCS TEAM from CSH Scenarios by subject # (102-117)



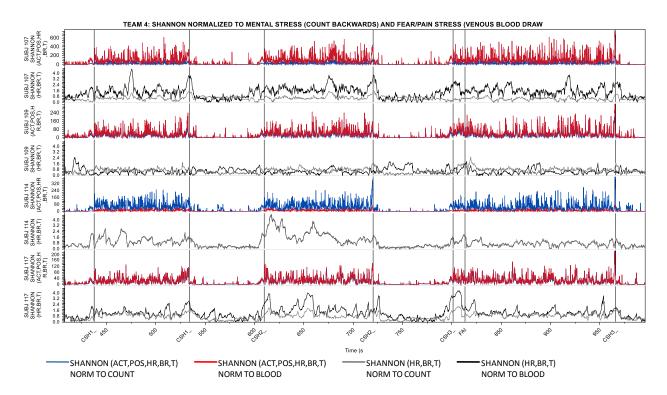




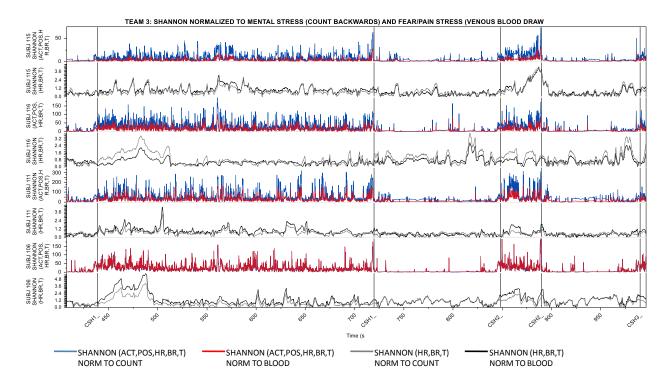


APPENDIX 7: SHANNON FILTERED DATA NORMALIZED TO MENTAL STRESS (Counting Backwards) AND FEAR/PAIN STRESS (venous blood draw)

TEAM4: Post-survey analysis identified Subject 114 as the primary leader, with Subjects 109 and 117 serving as secondary leaders. As is demonstrated below, Subject 114 has a much higher level of mental stress, as compared to fear/pain stress.

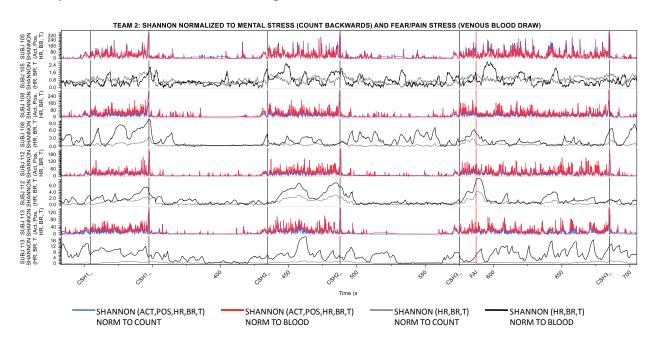


TEAM 3: Post-survey analysis identified leadership as shared between the team. In agreement with this, the graphs below indicate greater mental stress in Subjects 111, 115, and 116. Subject 106 appears to have a greater fear/pain response throughout the scenarios.



This trend does not appear perfect (as evidenced by TEAMS 1 & 2, shown below), but is worth exploring more.

TEAM 2: Post-survey analysis identified Subject 108 as the primary leader. It should be noted that Subject 105 nearly passed out during the venous blood draw (fear/pain basline), which is probably skewing his Shannon Norm to Blood (making it appear lower than it actually was) because of his extreme response to the blood draw.



TEAM 1: Post-survey analysis identified Subject 103 as the primary leader (with a minor role), and Subject 102 as the secondary leader (with a minor role). As is shown below, Subject 103 does have a higher mental stress (as compared to fear/pain stress), but Subject 102 does not. Subject 110 also has a higher mental stress as compared (as compared to fear/pain stress), but was not identified as a leader.

